Appl. No.: 10/573,212

Amdt. Dated February 1, 2008

Reply to Office Action of November 1, 2007

Amendments to the Claims:

1. (Currently Amended) A non-human <u>transgenic</u> animal non-responsive to endotoxin wherein a part or a whole of TRIF-related adaptor molecule (TRAM) genes on its chromosome is deleted, a function of expressing TRAM which is expressed in a wild-type is lacked, and responsiveness to a ligand recognized by TLR4 is specifically impaired.

- 2. (Currently Amended) The non-human <u>transgenic</u> animal non-responsive to endotoxin according to claim 1, which is responsive to PGN, R-848 and CpG ODN, and non-responsive to LPS.
- 3. (Currently Amended) The non-human <u>transgenic</u> animal non-responsive to endotoxin according to claim 1 or 2 wherein the non-human animal is a mouse.
- 4. (Currently Amended) A method for screening a substance promoting or suppressing a response to a ligand recognized by TLR4, wherein the response to the ligand recognized by TLR4 in a immunocyte derived from the non-human <u>transgenic</u> animal non-responsive to endotoxin according to <u>any one of claims 1 to 3 claim 1 or 2</u> is measured/estimated, with the use of the immunocyte, a test substance and the ligand recognized by TLR4.
- 5. (Currently Amended) A method for screening a substance promoting or suppressing a response to a ligand recognized by TLR4, wherein the response to the ligand recognized by TLR4 in the non-human <u>transgenic</u> animal according to <u>any one of claims 1 to 3 claim 1 or 2</u> is measured/estimated, with the use of the non-human animal, a test substance and the ligand recognized by TLR4.
- 6. (New) The non-human transgenic animal non-responsive to endotoxin according to claim 4 wherein the non-human animal is a mouse.

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7. (New) The non-human transgenic animal non-responsive to endotoxin according to claim 5 wherein the non-human animal is a mouse.

8. (New) An immunocyte from the non-human transgenic animal non-responsive to endotoxin according to claim 1 or 2.